INVENTOR SEARCH

 \Rightarrow d ibib abs 13 1-2

L3 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2003:927129 HCAPLUS Full-text Regulated attenuation of live vaccines to enhance cross

protective immunogenicity

INVENTOR(S): Curtiss, Roy Iii

PATENT ASSIGNEE(S): Washington University, USA

SOURCE: PCT Int. Appl. CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA	PATENT NO.					KIND		DATE		APPLICATION NO.					DATE			
WO	2003096812			A1		20031127		WO 2003-US11802					20030415					
	W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,	
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KΖ,	LC,	LK,	LR,	
		LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NΙ,	NO,	NΖ,	OM,	
		PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	ТJ,	TM,	TN,	TR,	TT,	
		TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW						
	RW:	GH,	GM,	ΚE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	ΑM,	ΑZ,	BY,	
		KG,	KΖ,	MD,	RU,	ТJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	
		FI,	FR,	GB,	GR,	HU,	IE,	ΙΤ,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,	
		BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	ΤG	
CA	CA 2526895				A1 20031127				CA 2003-2526895					20030415				
AU	AU 2003235457				A1 20031202				AU 2003-235457					20030415				
EP	1499	191			A1		2005	0126		EP 2	003-	7217	11		2	0030	415	
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙΤ,	LI,	LU,	NL,	SE,	MC,	PT,	
		IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	SK		
US	2006	0233	829		A1		2006	1019		US 2	005-	5116	16		2	0051	115	
RIORIT	ORITY APPLN. INFO.:									US 2002-372616P				P 20020415				
									,	US 2	002-	3736	26P		P 2	0020	418	
									,	WO 2	003-	US11	802	1	W 2	0030	415	
D 7	7 2			امما	۔ کی میں میں آم	د عدمات		e	مالامم		ما ما			سد مداد	م ام مہ	م کے ا		

AB A live attenuated derivative of a pathogenic bacterium intended for use as a vaccine.

OS.CITING REF COUNT: 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD

(2 CITINGS)

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1991:469825 HCAPLUS Full-text

DOCUMENT NUMBER: 115:69825

ORIGINAL REFERENCE NO.: 115:12050h,12051a

TITLE: Cross-protective Salmonella

vaccines using multiply mutant S. typhimurium

INVENTOR(S): Curtiss, Roy, III; Munson, Maryann

PATENT ASSIGNEE(S): Washington University, USA

SOURCE: PCT Int. Appl., 64 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA	PATENT NO.					KIND		DATE		APPLICATION NO.						DATE
WO	9106	 317			A1	_	1991	0516	W	10	1990-	 US65()3		-	19901102
	W:	ΑU,	CA,	JP												
	RW:	ΑT,	BE,	CH,	DE,	DK	, ES,	FR,	GB,	GF	R, IT,	LU,	NL,	SE		
CA	2072	633			A1		1991	0504	С	ĊΑ	1990-	20726	33			19901102
AU	9067	371			A		1991	0531	A	U	1990-	67371	_			19901102
EP	5006	99			A1		1992	0902	E	ΞP	1990-	9170	76			19901102
EP	P 500699				В1		1998	0610								
	R:	ΑT,	BE,	CH,	DE,	DK	, ES,	FR,	GB,	GF	R, IT,	LI,	NL,	SE		
JP	0550	4331			Т		1993	0708	J	Ρ	1990-	51588	38			19901102
AT	1670	61			Т		1998	0615	A	T	1990-	9170	76			19901102
PRIORITY APPLN. INFO.:									U	JS	1989-	43159	7		Α	19891103
									W	Ю	1990-	US650)3		Α	19901102

AB Attenuated Salmonella for use as live vaccines against Salmonella and other Gram-neg. bacteria are prepared The organisms are incapable of manufacturing the lipopolysaccharide involved in pathogenesis because of mutation in several genes involved in the synthesis of, or regulation of synthesis of, the lipopolysaccharide. Other genes involved in the regulation of pathogenesis-related genes are also inactivated. A S. typhimurium with the crp and cya genes deleted was prepared by transposon mutagenesis with Tn10. S. typhimurium carrying both deletions had an LD50 of >109 colony-forming units (CFU) in Leghorn chicks, vs. 2 + 104 - 2 + 105 for wild-types. Similar deletions of the phoP, fur, pmi, and gale genes were constructed. Some of the constructs prepared were found to confer cross-resistance to S. enteriditis and pathogenic Escherichia coli.

OS.CITING REF COUNT: 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (4 CITINGS)

3

RESULTS FROM SEARCHES IN CAPLUS, MEDLINE, BIOSIS, EMBASE, AND DRUGU

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=> d que stat 121
        182203 SEA FILE=HCAPLUS ABB=ON ?SALMONELLA? OR E(W)?COLI?
L6
         33841 SEA FILE=HCAPLUS ABB=ON L5 AND (?LIVE? OR ?ATTENUAT? OR
               ?DERIV?)
           744 SEA FILE=HCAPLUS ABB=ON L6 AND ?REGULAT?(5A)?EXPRES?
L7
           744 SEA FILE=HCAPLUS ABB=ON L7 AND (?REGULAT? OR ?PROMOT? OR FUR
L8
               OR ?INTEST? OR ?EXPRES?)
           189 SEA FILE=HCAPLUS ABB=ON L8 AND (?CARBOHYDRAT?(W)?ANTIGEN? OR
L9
                IN(W)?VIVO? OR ?CROSS?(W)?PROTECT? OR ?IMMUN?)
            21 SEA FILE=HCAPLUS ABB=ON L9 AND LPS
L11
L12
             1 SEA FILE=HCAPLUS ABB=ON L11 AND O(W)?ANTIGEN?
L14
             1 SEA FILE=HCAPLUS ABB=ON L9 AND PMI
L16
            22 SEA FILE=HCAPLUS ABB=ON L11 OR L12 OR L14
            18 SEA FILE=HCAPLUS ABB=ON L16 AND (PRD<20070418 OR PD<20070418)
L17
           114 SEA L17
L18
L19
           114 SEA L18 AND (LPS OR PMI)
L20
             3 SEA L19 AND O(W) ?ANTIGEN?
L21
            21 DUP REMOV L17 L20 (0 DUPLICATES REMOVED)
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=> d ibib abs 121 1-21

L21 ANSWER 1 OF 21 MEDLINE on STN

ACCESSION NUMBER: 2007442012 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 17629369

TITLE: Core-linked LPS expression of Shigella dysenteriae serotype 1 O-antigen in live Salmonella Typhi vaccine vector

Ty21a: preclinical evidence of immunogenicity and

protection.

Xu De Qi; Cisar John O; Osorio Manuel; Wai Tint T; Kopecko AUTHOR:

Dennis J

CORPORATE SOURCE: Laboratory of Enteric and Sexually Transmitted Diseases,

FDA-CBER, Bethesda, MD 20892, United States..

dennis.kopecko@fda.hhs.gov

Vaccine, (2007 Aug 14) Vol. 25, No. 33, pp. SOURCE: 6167-75. Electronic Publication: 2007-06-26.

Journal code: 8406899. ISSN: 0264-410X.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE) (RESEARCH SUPPORT, N.I.H., INTRAMURAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200712

Entered STN: 31 Jul 2007 ENTRY DATE:

> Last Updated on STN: 11 Dec 2007 Entered Medline: 6 Dec 2007

AΒ Shigella dysenteriae serotype 1 (S. dysenteriae 1) causes severe shigellosis that is typically associated with high mortality. Antibodies against Shigella serotype-specific O-polysaccharide (O-Ps) have been shown to be host protective. In this study, the rfb locus and the rfp gene with their cognate promoter regions were PCR-amplified from S. dysenteriae 1, cloned, and sequenced. Deletion analysis showed that eight rfb ORFs plus rfp are necessary for biosynthesis of this O-Ps. A tandemly-linked rfb-rfp gene cassette was cloned into low copy plasmid pGB2 to create pSd1. Avirulent Salmonella enterica serovar Typhi (S. Typhi) Ty21a harboring pSd1 synthesized

S. Typhi 9, 12 LPS as well as typical core-linked S. dysenteriae 1 LPS. Animal immunization studies showed that Ty21a (pSd1) induces protective immunity against high stringency challenge with virulent S. dysenteriae 1 strain 1617. These data further demonstrate the utility of S. Typhi Ty21a as a live, bacterial vaccine delivery system for heterologous O-antigens, supporting the promise of a bifunctional oral vaccine for prevention of shigellosis and typhoid fever.

L21 ANSWER 2 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2006:1056021 HCAPLUS Full-text

DOCUMENT NUMBER: 145:354218

TITLE: Down-regulation of key virulence factors makes the Salmonella enterica serovar Typhimurium rfaH mutant a promising live-

attenuated vaccine candidate

AUTHOR(S): Nagy, Gabor; Danino, Vittoria; Dobrindt, Ulrich;

Pallen, Mark; Chaudhuri, Roy; Emody, Levente; Hinton,

Jay C.; Hacker, Jorg

CORPORATE SOURCE: Department of Medical Microbiology and Immunology,

University of Pecs, Pecs, 7624, Hung.

SOURCE: Infection and Immunity (2006), 74(10),

5914-5925

CODEN: INFIBR; ISSN: 0019-9567
American Society for Microbiology

DOCUMENT TYPE: Journal LANGUAGE: English

PUBLISHER:

Mutants of Salmonella enterica serovar Typhimurium that lack the AB transcriptional regulator RfaH are efficient as live oral vaccines against salmonellosis in mice. The authors show that the attenuation of the vaccine candidate strain is associated with reduced net growth in epithelial and macrophage cells. To identify the relevant RfaH-dependent genes, the RfaH regulon was determined with S. enterica serovars Enteritidis and Typhimurium using whole-genome Salmonella microarrays. As well as impacting the expression of genes involved in lipopolysaccharide (LPS) core and O- antigen synthesis, the loss of RfaH results in a marked down- regulation of SPI-4 genes, the flagellum/chemotaxis system, and type III secretion system 1. However, a proportion of these effects could have been the indirect consequence of the altered expression of genes required for LPS biosynthesis. Direct and indirect effects of the rfaH mutation were dissociated by genome-wide transcriptional profiling of a structural deep-rough %PS mutant (waaG). The authors show that truncation of LPS itself is responsible for the decreased intracellular yield observed for Δ rfaH strains. LPS mutants do not differ in replication ability; rather, they show increased susceptibility to antimicrobial peptides in the intracellular milieu. Evidence that deletion of rfaH, as well as some other genes involved in LPS biosynthesis, results in enhanced invasion of various mammalian cells is shown. Exposure of common minor antigens in the absence of serovar-specific antigens might be responsible for the observed cross-reactive nature of the elicited immune response upon vaccination. Increased invasiveness of the Salmonella rfaH mutant into antigen-presenting cells, combined with increased intracellular killing and the potential for raising a cross-protective immune response, renders the rfaH mutant an ideal vaccine candidate. OS.CITING REF COUNT: 17 THERE ARE 17 CAPLUS RECORDS THAT CITE THIS

RECORD (17 CITINGS)

REFERENCE COUNT: 62 THERE ARE 62 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 3 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2006:156747 HCAPLUS $\underline{\text{Full-text}}$

DOCUMENT NUMBER: 145:410249

TITLE: Immunostimulatory CpG-oligodeoxynucleotides

(CpG-ODN) induce early hepatic injury, but provide a late window for protection against endotoxin-mediated

liver damage

AUTHOR(S): Slotta, Jan E.; Scheuer, Claudia; Menger, Michael D.;

Vollmar, Brigitte

CORPORATE SOURCE: Institute for Clinical and Experimental Surgery,

University of Saarland, Homburg/Saar, 66421, Germany

SOURCE: Journal of Hepatology (2006), 44(3), 576-585

CODEN: JOHEEC; ISSN: 0168-8278

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

An impaired immunol, response to infection has been recognized as a major AB defect in the pathogenesis of sepsis and multi-organ failure. Sepsis-associated liver dysfunction and damage are main determinants for the course of the disease. CpG-motif-containing DNA-sequences (CpG-ODN) were previously shown to confer protection in models of infection by stimulating both innate and specific immune responses. Herein, the authors studied the effect of CpG-ODN in lipopolysaccharide (LPS)-associated hepatotoxicity. Sprague Dawley rats pre-treated at day 6 with either CpG-ODN or inert DNA were challenged with E. coli LPS and subsequently studied for liver injury at 6 and 16 h using in vivo fluorescence microscopy and immunohistochem. Western blot protein anal. served for assessment of expression of TLR4, TNF receptor-associated factor 6 (TRAF6), NFKB, and caspase-3. To evaluate CpG-ODN effects during non-septic conditions, addnl. animals were solely exposed to CpG-ODN and studied after 1 and 6 days. CpG-ODN application induced marked hepatic microcirculatory deterioration and liver dysfunction at day 1, however, with almost complete recovery to normal at day 6. Interestingly, CpG-ODN pre-treatment decreased LPS -induced leukocyte-endothelial cell interaction, sinusoidal perfusion failure, and caspase-3-dependent apoptotic cell death. Although Kupffer cell phagocytic activity was not affected, CpG-ODN pre-treatment in LPS-challenged animals attenuated hepatic protein expression of TRAF6 and NFKB and increased TLR4 by almost 100%. Thus, CpG-containing DNA-sequences induce early hepatic injury, but mediate long-term protection against LPS hepatotoxicity. The mechanism of protection is based on the induction of cross-tolerance, probably via inhibition of the downstream $TRAF6-NF\kappa B$ signaling pathway and upregulation of the TLR4 surface receptor. OS.CITING REF COUNT: 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD

(4 CITINGS)

REFERENCE COUNT: 62 THERE ARE 62 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 4 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2005:427114 HCAPLUS Full-text

DOCUMENT NUMBER: 143:58451

TITLE: Insect renal tubules constitute a cell-autonomous

immune system that protects the organism

against bacterial infection

AUTHOR(S): McGettigan, J.; McLennan, R. K. J.; Broderick, K. E.;

Kean, L.; Allan, A. K.; Cabrero, P.; Regulski, M. R.; Pollock, V. P.; Gould, G. W.; Davies, S.-A.; Dow, J.

A. T.

CORPORATE SOURCE: Division of Molecular Genetics, Institute of

Biomedical and Life Sciences, University of Glasgow,

Glasgow, G12 8QQ, UK

SOURCE: Insect Biochemistry and Molecular Biology (

2005), 35(7), 741-754

CODEN: IBMBES; ISSN: 0965-1748

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

Innate immunity is a widespread and important defense against microbial attack, which in insects is thought to originate mainly in the fat body. Here we demonstrate that the fluid-transporting Malpighian (renal) tubule of Drosophila melanogaster constitutes an autonomous immune-sensing tissue utilizing the nitric oxide (NO) signaling pathway. Reverse transcriptase PCR (RT-PCR) shows that tubules express those genes encoding components of the Imd pathway. Furthermore, isolated tubules bind and respond to lipopolysaccharide (LPS), by uprequiating anti-microbial peptide (diptericin) gene expression and increased bacterial killing. Excised, LPS-challenged tubules, as well as tubules from LPS-infected flies, display increased NO synthase (NOS) activity upon immune challenge. Targetted expression of a Drosophila NOS (dNOS) transgene to only principal cells of the tubule main segment using the GAL4/UAS system increases diptericin expression . In live flies, such targeted over-expression of dNOS to tubule principal cells confers increased survival of the whole animal upon E. coli challenge. Thus, we describe a novel role of Malpighian tubules in immune sensing and insect survival.

OS.CITING REF COUNT: 26 THERE ARE 26 CAPLUS RECORDS THAT CITE THIS

RECORD (26 CITINGS)

REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 5 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2005:674510 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 143:246658

TITLE: Activation of murine dendritic cells and macrophages

induced by Salmonella enterica serovar

Typhimurium

AUTHOR(S): Kalupahana, Ruwani Sagarika; Mastroeni, Pietro;

Maskell, Duncan; Blacklaws, Barbara Ann

CORPORATE SOURCE: Centre for Veterinary Science, Department of

Veterinary Medicine, University of Cambridge,

Cambridge, UK

SOURCE: Immunology (2005), 115(4), 462-472

CODEN: IMMUAM; ISSN: 0019-2805

PUBLISHER: Blackwell Publishing Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

Macrophages and dendritic cells (DCs) are antigen-presenting cells (APCs), AΒ and the direct involvement of both cell types in the immune response to Salmonella has been identified. In this study the authors analyzed the phenotypic and functional changes that take place in murine macrophages and DCs in response to live and heat-killed Salmonella enterica serovar Typhimurium. Both types of cell secreted proinflammatory cytokines and nitric oxide (NO) in response to live and heat-killed salmonellae. Bacterial stimulation also resulted in up-regulation of costimulatory mols. on macrophages and DCs. The expression of major histocompatibility complex (MHC) class II mols. by macrophages and DCs was differentially regulated by interferon (IFN)-y and salmonellae. Live and heat-killed salmonellae as well as lipopolysaccharide (LPS) inhibited the up-regulation of MHC class II expression induced by IFN-y on macrophages but not on DCs. Macrophages as well as DCs presented Salmonella -derived antigen to CD4 T cells, although DCs were much more efficient than macrophages at stimulating CD4 T-cell cytokine release. Macrophages are effective in the uptake and killing of bacteria while DCs specialize in antigen presentation. This study showed that the viability of salmonellae was not essential for activation of APCs but, unlike live bacteria, prolonged contact with heat-killed bacteria was necessary to obtain maximal expression of the activation markers studied.

OS.CITING REF COUNT: 6 THERE ARE 6 CAPLUS RECORDS THAT CITE THIS RECORD (6 CITINGS)

REFERENCE COUNT: 68 THERE ARE 68 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 6 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2005:864561 HCAPLUS $\underline{\text{Full-text}}$

DOCUMENT NUMBER: 143:345201

TITLE: Role of nuclear transcription factor- κ B in

endotoxin-induced shock in rats

AUTHOR(S): Wang, Jin; Yang, Guangtian

CORPORATE SOURCE: Emergency Department, Tongji Hospital, Tongji Medical

College, Huazhong University of Science and Technology, Wuhan, 430030, Peop. Rep. China Journal of Huazhong University of Science and Technology, Medical Sciences (2005), 25(2),

174-177

CODEN: JHUSAW; ISSN: 1672-0733

PUBLISHER: Huazhong University of Science and Technology

DOCUMENT TYPE: Journal LANGUAGE: English

SOURCE:

To investigate the role of NF- κ B in endotoxic shock in rats, the shock was induced by i.v. infusion of E. coli lipopolysaccharide (LPS). At 1 h, 2 h, 4 h, and 6 h after LPS injection, the activation of NF- κ B in blood mononuclear cells and the content of TNF-lpha and IL-6 in plasma were detected by enzymelinked immunoadsorbent assay (ELISA). The level of mean arterial pressure (MAP) and the histopathol. changes of lung and liver were also observed The activation of NF- κ B in mononuclear cells increased 1 h after LPS injection and reached its peak 2 h after the injection, and its level was higher than in the normal group. The level of TNF- α was increased 1 h after the infusion and peaked 2 h after the injection, and its level was higher than that of normal group after LPS infusion. The content of IL-6 increased gradually with time, and the IL-6 level was higher than in the normal group after LFS injection. MAP decreased gradually with time and its level was lower than in the normal group after LPS injection. Pathol. examination showed that endotoxic shock could cause pulmonary alveolar hemorrhage, edema, and infiltration of inflammatory cells into lung, and congestion, edema, capillary dilation, and inflammatory cell infiltration in liver tissue. Thus, NF-KB can up-regulate the expression of TNF- α and IL-6 in plasma and therefore plays an important role in endotoxin-induced shock in rats.

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 7 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2005:581386 HCAPLUS Full-text

DOCUMENT NUMBER: 143:324458

TITLE: Cytokine gene expression profiles of bovine

dendritic cells after interaction with Mycobacterium avium ssp. paratuberculosis (M.a.p.), Escherichia coli

(E. coli) or recombinant M.a.p.

heat shock protein 70

AUTHOR(S): Langelaar, Merel F. M.; Weber, Corinna N.; Overdijk,

Marije B.; Mueller, Kerstin E.; Koets, Ad P.; Rutten,

Victor P. M. G.

CORPORATE SOURCE: Division of Immunology, Department of Infectious

Diseases and Immunology, Faculty of Veterinary

Medicine, Utrecht University, Utrecht, 3508 TD, Neth.

SOURCE: Veterinary Immunology and Immunopathology (

2005), 107(1-2), 153-161

CODEN: VIIMDS; ISSN: 0165-2427

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

Mycobacterium avium paratuberculosis (M.a.p.) resides and replicates in macrophages. Many of the of immune mechanisms aiding M.a.p. survival in the host's cells are known. However, little is known about interactions of M.a.p. with dendritic cells (DC). As DC are important for the induction of protective immunity against infectious diseases, we investigated the interaction of M.a.p. with these cells. Quant. real-time PCR (RT-PCR) was used to analyze differential expression of cytokine genes after 6 h and 24 h of incubation by immature DC that phagocytosed either M.a.p. or Escherichia coli (& . coli). We hypothesized that phagocytosis of M. coli would induce pro-inflammatory cytokines due to abundant presence of lipopolysaccharide (LPS) and that the cytokine expression profile induced by phagocytosis of live M.a.p. would differ. In addition we hypothesized that incubation of immature DC with rHsp70, an immunodominant antigen of M.a.p., would induce a similar profile of cytokine gene expression as phagocytosis of intact M.a.p. However, phagocytosis of both E. coli and M.a.p. resulted in a cytokine gene expression pattern representative of a (pro-)inflammatory reaction, dominated by strong induction of IL-12 gene expression, that was higher after 24 h than after 6 h of incubation, although the response to M.a.p. was less vigorous than to E. coli. Incubation with rHsp70 resulted in a more inhibitory type of cytokine gene expression, with delayed IL-12 gene expression and downregulation of the genes for IL-1B and IL-6 after 24 h of incubation. We conclude that bovine DC produce an immuno-stimulatory, anti-mycobacterial response to infection with M.a.p., while Hsp70 potentially contributes to pathogen virulence by allowing the bacteria to invade the host cell.

OS.CITING REF COUNT: 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD

(4 CITINGS)

REFERENCE COUNT: 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 8 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:759609 HCAPLUS Full-text

DOCUMENT NUMBER: 141:294659

TITLE: Expression of inflammatory and septic genes

to identify antiinflammatory and antiseptic peptides

for therapeutic use

INVENTOR(S): Hancock, Robert E. W.; Finlay, B. Brett; Scott,

Monisha Gough; Bowdish, Dawn; Rosenberger, Carrie

Melissa; Powers, Jon-Paul Steven

PATENT ASSIGNEE(S): Can.

SOURCE: U.S. Pat. Appl. Publ., 93 pp., Cont.-in-part of U.S.

Pat. Appl. 2004 1,803.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE			
US 20040180038	A1	20040916	US 2003-661471	20030912 <			
US 20040001803	A1	20040101	US 2002-308905	20021202 <			
US 7507787	В2	20090324					
CN 101215601	A	20080709	CN 2007-10168028	20021202 <			
NZ 563261	A	20080829	NZ 2002-563261	20021202 <			
AU 2004271668	A1	20050324	AU 2004-271668	20040910 <			
CA 2538069	A1	20050324	CA 2004-2538069	20040910 <			
WO 2005025607	A1	20050324	WO 2004-CA1602	20040910 <			
W: AE, AG, AI	, AM, AT,	AU, AZ, B	A, BB, BG, BR, BW, BY,	BZ, CA, CH,			
CN, CO, CF	, CU, CZ,	DE, DK, DI	M, DZ, EC, EE, EG, ES,	FI, GB, GD,			
GE, GH, GN	HR, HU,	ID, IL, II	N, IS, JP, KE, KG, KP,	KR, KZ, LC,			

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LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
              NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
              TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW
          RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
              AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
              EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
              SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
              SN, TD, TG
                                   20060614
                                                 EP 2004-761766
                                                                           20040910 <--
     EP 1667707
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              IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK
                                              CN 2004-80031833
     CN 1901933
                                   20070124
                                                                           20040910 <--
                            Α
     JP 2007505048
                             Τ
                                   20070308
                                                 JP 2006-525585
                                                                           20040910 <--
                                                 ZA 2006-2754
     ZA 2006002754
                                   20070627
                                                                           20040910 <--
                           Α
                           A1 20070816 US 2005-241882
     US 20070190533
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A 20070817 IN 2006-DN1886
     MX 2006002828
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     IN 2006DN01886
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     US 20070134261 A1 20070614 US 2006-400411 KR 2007033314 A 20070326 KR 2006-706853 AU 2007201885 A1 20070517 AU 2007-201885
                                                                           20060407 <--
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                                                 US 2001-336632P P 20011203 <--

US 2002-308905 A2 20021202 <--

AU 2002-365675 A3 20021202 <--
PRIORITY APPLN. INFO.:
                                                 AU 2002-365675
                                                                      A3 20021202 <--
                                                 CN 2002-827327
NZ 2002-533721
US 2003-661471
                                                                      A3 20021202 <--
                                                                      A3 20021202 <--
                                                                       A 20030912 <--
                                                 WO 2004-CA1602
US 2005-241882
                                                                       W 20040910 <--
                                                                       A2 20050929 <--
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AB A method of identifying a polynucleotide or pattern of polynucleotides regulated by one or more sepsis or inflammatory inducing agents and inhibited by a peptide is described. A method of identifying a pattern of polynucleotide expression for inhibition of an inflammatory or septic response. The method includes contacting human epithelial cells with LPS, LTA, CpG DNA and/or intact microbe or microbial components in the presence or absence of a cationic peptide; detecting a pattern of polynucleotide expression for the cells in the presence and absence of the peptide, wherein the pattern in the presence of the peptide represents inhibition of an inflammatory or septic response. Also included are compds. and agents identified by the methods of the invention. In another aspect, the invention provides methods and compds. for enhancing innate immunity in a subject. OS.CITING REF COUNT: 7 THERE ARE 7 CAPLUS RECORDS THAT CITE THIS RECORD

(7 CITINGS)

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L21 ANSWER 9 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN
ACCESSION NUMBER:
                        2005:16780 HCAPLUS Full-text
DOCUMENT NUMBER:
                        142:217022
TITLE:
                        Over-expression of AIF-1 in liver
                        allografts and peripheral blood correlates with acute
                        rejection after transplantation in rats
AUTHOR(S):
                        Nagakawa, Yuichi; Nomoto, Shuji; Kato, Yukihiko;
                        Montgomery, Robert A.; Williams, George Melville;
                        Klein, Andrew S.; Sun, Zhaoli
CORPORATE SOURCE:
                        Department of Surgery, Johns Hopkins University School
                        of Medicine, Baltimore, MD, USA
SOURCE:
                        American Journal of Transplantation (2004),
                        4(12), 1949-1957
                        CODEN: AJTMBR; ISSN: 1600-6135
                       Blackwell Publishing Ltd.
PUBLISHER:
DOCUMENT TYPE:
                        Journal
                        English
LANGUAGE:
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Early and accurate detection of acute cellular rejection (ACR) is important AB in the management of liver allograft recipients. We hypothesized that expression of allograft inflammatory factor (AIF)-1 would be associated with liver allograft rejection as previous studies have shown that a relationship exists between kidney and heart transplantation. Indeed using rat orthotopic transplant models we found that the expression of allograft inflammatory factor-1 (AIF-1) can be detected in both allograft and peripheral blood leukocytes with peak levels detected 7 days following liver transplantation. Interestingly, AIF-1 expression increased 2-fold in acutely rejecting liver allografts compared to chronically accepted livers on days 5, 7 and 10 after transplantation. AIF-1 expression in peripheral blood leukocytes was also significantly greater in the rejection model than in the acceptance model. Flow cytometric anal. of peripheral blood leukocytes demonstrated that AIF-1 was expressed in ED2-pos. cells, a marker for Kupffer cells. In vitro studies showed that AIF-1 expression in Kupffer cells was upregulated by coculture with Th1 cytokines. However, neither LPS nor Escherichia colt (E. coli) administration had an affect on AIF-1 expression. These data indicate that high levels of AIF-1 expression reflect aggressive liver allograft rejection and suggest a role for monitoring AIF-1 in peripheral blood leukocytes as 7 a monitor for increased immunosuppression. OS.CITING REF COUNT: THERE ARE 7 CAPLUS RECORDS THAT CITE THIS RECORD

(7 CITINGS)

REFERENCE COUNT: 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 10 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:558899 HCAPLUS Full-text

DOCUMENT NUMBER: 141:258441

TITLE: Implication of CpG-ODN and reactive oxygen species in

the inhibition of intracellular growth of Salmonalla typhimurium in hepatocytes

AUTHOR(S): Sanchez-Campillo, Maria; Chicano, Antonio; Torio,

Alberto; Martin-Orozco, Elena; Gamiz, Pilar;

Hernandez-Caselles, Trinidad; Garcia-Penarrubia, Pilar CORPORATE SOURCE: Department of Biochemistry and Molecular Biology B and

Immunology, School of Medicine, University of Murcia,

Muncia 20100 Chain

Murcia, 30100, Spain

SOURCE: Microbes and Infection (2004), 6(9), 813-820

CODEN: MCINFS; ISSN: 1286-4579

PUBLISHER: Elsevier Science B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

Bacterial DNA acts as an alert signal for eukaryotic cells through immunostimulatory CpG motifs. These sequences have therapeutic properties promoting protective immune TH1 responses and are recognized by a membrane protein belonging to the Toll-like receptor (TLR) family, named TLR-9. The aim of this study was to test the capability of murine hepatocytes to sense bacterial DNA and to develop antibacterial mechanisms against Salmonella typhimurium. The authors show that hepatocyte cell lines and mRNA exts. from murine liver constitutively express TLR-9, which is down-regulated by LPS and the mix of IFNy, IL-1 β and LPS. Also, the authors have found that hepatocyte cell lines can sense the presence of bacterial DNA and respond to it by increasing the pool of intracellular peroxides. This results in inhibition of intracellular growth of S. typhimurium when infected cells were incubated in the presence of CpG synthetic oligonucleotides (CpG-ODN). Empression of hepatocyte Mn-SOD is also induced by stimulation with CpGoligodeoxynucleotides, LPS, and the mix of IFNy, IL-1 β and LPS. These results reinforce the prominent role of hepatocytes as a microbial product-responsive cell and the capabilities of CpG-ODN sequences as potent inducers of the innate immune response through the activation of a broad range of cell types.

OS.CITING REF COUNT: 5 THERE ARE 5 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS)

REFERENCE COUNT: 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 11 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2005:1164833 HCAPLUS Full-text

DOCUMENT NUMBER: 145:25990

TITLE: The context of host encounter with the flagellin protein, FliC, determines the direction of the T

halper response

helper response

AUTHOR(S): Cunningham, A. F.; Toellner, K. M.; Serre, K.; Mohr,

E.; Khan, M.; Ball, J.; MacLennan, I. C. M.

CORPORATE SOURCE: MRC Centre for Immune Regulation, University of

Birmingham, Birmingham, UK

SOURCE: Immunology 2004, [12th International Congress of

Immunology and 4th Annual Conference of FOCIS], Montreal, QC, Canada, July 18-23, 2004 (2004)

), E718C4131/1-E718C4131/4. Monduzzi Editore: Bologna, Italy.

CODEN: 69HJYL; ISBN: 88-7587-070-5

DOCUMENT TYPE: Conference; (computer optical disk)

LANGUAGE: English

Features of the Th1 or Th2 phenotype start to develop during CD4 T cell priming, when dendritic cells (DC) present antigen to T cells. Divergent polarization of mouse Th responses to the same protein is exemplified by responses to the bacterial flagellar protein, FliC. This is strictly Th2 following immunization with soluble, recombinant (r) FliC or isolated bacterial flagellae. Conversely, when presented in its native context as flagellated live Salmonella a Th1 response is induced. The polarization of a native - non-transgenic - anti-FliC response is shown in situ by the way primed CD4 T cells induce class switching in FliC-specific B cells. The presence of FliC-specific plasma cells producing IgG2a (IgG2c in C57B1/6 mice) reflects Th1 activity, while switching to IgG1 indicates Th2 activity. Thus soluble rFliC and polymerized FliC isolated from the surface of Salmonella predominantly induce IgG1 switching. Conversely, against FliC surface-bound to live, flagellated Salmonella, Th1-reflecting IgG2c is predominantly induced. Thus the intrinsic, innate immunostimulatory properties of the FliC mol. are insufficient to drive a Th1 immune response. Addnl., soluble rFliC is shown, like LPS, to be a potent uprequiator of CD40 and CD86 on CD11c+ DC. By contrast LPS, but not rFliC upregulates the expression of CD40 and CD86 on B cells.

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 12 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:653792 HCAPLUS $\underline{\text{Full-text}}$

DOCUMENT NUMBER: 141:364779

TITLE: Differential regulation of membrane CD14

expression and endotoxin-tolerance in alveolar

macrophages

AUTHOR(S): Lin, Shu-Min; Frevert, Charles W.; Kajikawa, Osamu;

Wurfel, Mark M.; Ballman, Kimberly; Mongovin, Stephen;

Wong, Venus A.; Selk, Amy; Martin, Thomas R.

CORPORATE SOURCE: Pulmonary Research Laboratories at the VA Puget Sound

Medical Center, and the Division of Pulmonary/Critical Care Medicine, Department of Medicine, University of Washington School of Medicine, Seattle, WA, USA

SOURCE: American Journal of Respiratory Cell and Molecular

Biology (2004), 31(2, Pt. 1), 162-170

CODEN: AJRBEL; ISSN: 1044-1549

PUBLISHER: American Thoracic Society

DOCUMENT TYPE: Journal

LANGUAGE: English

CD14 is important in the clearance of bacterial pathogens from lungs. However, the mechanisms that regulate the expression of membrane CD14 (mCD14) on alveolar macrophages (AM) have not been studied in detail. This study examines the regulation of mCD14 on AM exposed to Escherichia coli in vivo and in vitro, and explores the consequences of changes in mCD14 expression. The expression of mCD14 was decreased on AM exposed to E. coli in vivo and AM incubated with lipopolysaccharide (LPS) or E. coli in vitro. Polymyxin B abolished LPS effects, but only partially blocked the effects of % . coli. Blockade of extracellular signal-regulated kinase pathways attenuated LPS and E. coli-induced decrease in mCD14 expression. Inhibition of proteases abrogated the LPS-induced decrease in mCD14 expression on AM and the release of sCD14 into the supernatants, but did not affect the response to Σ . colú. The production of tumor necrosis factor- α in response to a second challenge with Staphylococcus aureus or zymosan was decreased in AM after incubation with %. coli but not LPS. These studies show that distinct mechanisms regulate the expression of mCD14 and the induction of endotoxin tolerance in AM, and suggest that AM function is impaired at sites of bacterial infection. OS.CITING REF COUNT: 14 THERE ARE 14 CAPLUS RECORDS THAT CITE

RECORD (14 CITINGS)

REFERENCE COUNT: 46 THERE ARE 46 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 13 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:636885 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 141:343958

TITLE: Cloning and expression of porcine

adiponectin, and its relationship to adiposity,

lipogenesis and the acute phase response

AUTHOR(S): Jacobi, S. K.; Ajuwon, K. M.; Weber, T. E.; Kuske, J.

L.; Dyer, C. J.; Spurlock, M. E.

CORPORATE SOURCE: Department of Animal Sciences, Comparative Medicine

Program, Purdue University, West Lafayette, IN, USA

SOURCE: Journal of Endocrinology (2004), 182(1),

133-144

CODEN: JOENAK; ISSN: 0022-0795

PUBLISHER: Society for Endocrinology

DOCUMENT TYPE: Journal LANGUAGE: English

Adiponectin is an adipocyte-derived hormone that has been implicated recently in the regulation of inflammation in immunocytes, and in lipid metabolism and qlucose homeostasis in liver, skeletal muscle and adipocytes. However, information in non-rodent models is limited. We have cloned and sequenced the porcine adiponectin open reading frame and evaluated the regulation of adiponectin in vivo following lipopolysaccharide (LPS) or E. coli administration. The porcine sequence shares approx. 88, 86, 85 and 83% homol. with the dog, human, cow and mouse adiponectin resp., and 79-83% similarity with dog, human, cow and mouse proteins at the amino acid level, based on the translated porcine sequence and GenBank submissions for the other species. Relative serum adiponectin concns. were not altered in pigs infused with E. coli, and mRNA expression in adipose tissue was not responsive to LPS. However, anal. of serum from very lean vs. a substantially fatter genotype of pig indicated that relative circulating adiponectin concns. are higher (P<0.01) in the lean pigs than in the fatter genotype, and that the difference is established relatively early in the growth curve. Also, incubating pig adipocytes for 6 h with recombinant pig adiponectin resulted in an approx. 30% reduction (P < 0.05) in lipogenesis compared with adipocytes under basal conditions and with those incubated in the presence of insulin. This is the first report in any species that adiponectin antagonizes the incorporation of glucose carbon into lipid in the adipocyte, and provides addnl. evidence that adiponectin acts as an autocrine regulatory factor to regulate energy metabolism

OS.CITING REF COUNT: 20 THERE ARE 20 CAPLUS RECORDS THAT CITE THIS

RECORD (20 CITINGS)

REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 14 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2003:265342 HCAPLUS Full-text

DOCUMENT NUMBER: 138:319537

TITLE: Monocyte-derived RANTES is intrinsically

> elevated in periodontal disease while MCP-1 levels are related to inflammation and are inversely correlated

with IL-12 levels

Fokkema, S. J.; Loos, B. G.; Van Der Velden, U. AUTHOR(S): CORPORATE SOURCE: Department of Periodontology, Academic Centre for

Dentistry Amsterdam (ACTA), Neth.

SOURCE: Clinical and Experimental Immunology (2003),

131(3), 477-483

CODEN: CEXIAL; ISSN: 0009-9104

Blackwell Publishing Ltd. PUBLISHER:

DOCUMENT TYPE: Journal LANGUAGE: English

Bacteria colonizing tooth surfaces are essential in the induction of an inflammatory response in the periodontal tissues, but do not cause periodontitis in everyone, implicating differences in the host immune response. These possible differences were studied using lipopolysaccharide (LPS)-stimulated whole blood cell cultures (WBCC), which revealed a down regulation of monocyte derived interleukin-12 (IL-12p70) in untreated periodontitis patients and an up regulation after therapy. IL-12p70 is a crucial factor in the differentiation of Th1 cell responses. Since CC chemokines are able to influence the T cell differentiation via cytokine secretion in antigen-presenting cells, the production of CC chemokines in periodontitis was evaluated. Therefore WBCC were stimulated with LPS from Escherichia coli for 18 h and the levels of IL-12p70 and CC chemokines were measured in the supernatants by ELISA. Untreated periodontitis patients released 2 fold more RANTES (regulated on activation normal T cell expressed and secreted) (P = 0.01) and lower levels of IL-12p70 in comparison to controls (P < 0.05). A trend towards higher levels of macrophage chemoattractant protein-1 (MCP-1) (P = 0.07) was also seen in untreated periodontitis patients; while similar levels of monocyte dexived chemokine (MDC) and macrophage inflammatory proteins- 1α and -1β (MIP- 1α and -1β) were found. After periodontal therapy no changes were seen with regard to MDC, MIP-1 α , MIP-1 β and RANTES, whereas the MCP-1 levels decreased (P < 0.05) and the IL-12p70 levels strongly increased (P < 0.01). The data showed a consistent inverse correlation between the levels of MCP-1 and IL-12p70, and their proportional changes after therapy correlated with the clin. inflammatory response after therapy. This indicates that the disease state requiates the release of IL-12p70 and MCP-1 in & . coli LPS-stimulated WBCC. In contrast, the persistent augmented levels of RANTES after therapy are suggestive for an intrinsic behavior. THERE ARE 9 CAPLUS RECORDS THAT CITE THIS RECORD 9

OS.CITING REF COUNT:

(9 CITINGS)

REFERENCE COUNT: THERE ARE 43 CITED REFERENCES AVAILABLE FOR THIS 43 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 15 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2002:554529 HCAPLUS Full-text

DOCUMENT NUMBER: 137:139106

Bacterium-induced CXCL10 secretion by osteoblasts can TITLE:

be mediated in part through Toll-like receptor 4

Gasper, Nancy A.; Petty, Cynthia C.; Schrum, Laura W.; AUTHOR(S):

Marriott, Ian; Bost, Kenneth L.

CORPORATE SOURCE: Department of Biology, University of North Carolina at

Charlotte, Charlotte, NC, 28223, USA SOURCE:

Infection and Immunity (2002), 70(8),

4075-4082

CODEN: INFIBR; ISSN: 0019-9567 PUBLISHER: American Society for Microbiology

DOCUMENT TYPE: Journal LANGUAGE: English

Two common pathogens known to cause bone infection, Salmonella and Staphylococcus aureus, were investigated to determine their abilities to induce chemokine expression in cultured mouse and human osteoblasts. While these cells are responsible for bone formation, the authors were surprised to find that they could respond to bacterial infection by upregulating expression of the chemokine CXCL10 (IP-10). However, there were significant differences in the abilities of the gram-neg. bacterium Salmonella and the gram-pos. bacterium S. aureus to induce empression of CXCL10. Reverse transcription-PCR and ELISA analyses showed high levels of Salmonella-induced CXCL10 mRNA and protein expression, resp., whereas the osteoblast response to S. aureus was significantly less. Consistent with these findings, Salmonella-derived lipopolysaccharide (LPS), but not S. aureus-derived peptidoglycan, could induce expression of CXCL10. An antibody against Toll-like receptor 4 (TLR4) could block the LPS-induced CXCL10 production, demonstrating the functional expression of TLR4 by osteoblasts. Despite the inducible nature of TLR2 mRNA expression by bacterium-infected osteoblasts, peptidoglycan failed to stimulate CXCL10 secretion. Immunofluorescent staining of bacterium-infected calvaria (i.e., skull bone) demonstrated the presence of CXCL10 in osteoblasts. The fact that osteoblasts did not express CXCR3 mRNA, whereas T lymphocytes can express high levels of this receptor, suggests that osteoblast-derived CXCL10 may recruit T lymphocytes to the sites of bone infections.

OS.CITING REF COUNT: 30 THERE ARE 30 CAPLUS RECORDS THAT CITE THIS

RECORD (30 CITINGS)

THERE ARE 66 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 66 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 16 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2002:596518 HCAPLUS Full-text

DOCUMENT NUMBER: 137:306451

Molecular cloning, gene organization, and TITLE:

expression of mouse Mpi encoding

phosphomannose isomerase

Davis, Joseph A.; Wu, Xiao-Hua; Wang, Ling; DeRossi, AUTHOR(S):

Charles; Westphal, Vibeke; Wu, Rongrong; Alton, Gordon; Srikrishna, Geetha; Freeze, Hudson H.

Glycobiology Program, The Burnham Institute, La Jolla, CORPORATE SOURCE:

CA, 92037, USA

SOURCE: Glycobiology (2002), 12(7), 435-442

CODEN: GLYCE3; ISSN: 0959-6658

PUBLISHER: Oxford University Press

DOCUMENT TYPE: Journal LANGUAGE: English

Phosphomannose isomerase (FMI) interconverts fructose-6-P (Fru-6-P) and mannose-6-P (Man-6-P), linking energy metabolism to protein glycosylation. The authors have cloned the mouse Mpi cDNA, analyzed its genomic organization, and studied the expression in different tissues. The Mpi gene has eight exons covering 7.2 kb. The structure and intron-exon boundaries are essentially the same as its human ortholog with 85% amino acid identity. Mpi is alternatively spliced at the 3' end, resulting in three messages with different 3'-untranslated regions. Mpi empression is regulated at both the transcription and translation levels, with the highest expression level in testis. Rabbit antibodies prepared against mouse PMI expressed in E. coli recognize a single 47-kDa band. Immunohistochem. of mouse tissues shows general cytosolic staining in all cells. In testis, staining is intense in round spermatids and residual bodies, moderate in pachytene

spermatocytes, and weak in spermatogonia and spermatozoa. In contrast, northern blot anal. shows comparable transcripts of 1.8 and 1.6 kb in pachytene spermatocytes and round spermatids, suggesting delayed translation of PMI. The stage-specific expression of PMI in testis may be important for KDN synthesis, which requires Man-6-P, or it may be needed to ensure sufficient glycosylation precursors in cells that do not utilize glucose and instead rely on lactate and pyruvate. OS.CITING REF COUNT: 7 THERE ARE 7 CAPLUS RECORDS THAT CITE THIS RECORD

(7 CITINGS)

REFERENCE COUNT: 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 17 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1999:662054 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 131:335603

TITLE: The intestinal microflora regulates

cytokine production positively in spleenderived macrophages but negatively in bone

marrow-derived macrophages

AUTHOR(S): Nicaise, Pascale; Gleizes, Aude; Sandre, Catherine;

Kergot, Roseline; Lebrec, Herve; Forestier, Francoise;

Labarre, Colette

CORPORATE SOURCE: Department de Microbiologie et Immunologie, Unite

Associee INRA Ecologie Microbienne du Tube Digestif et

Sante, Paris, Fr.

SOURCE: European Cytokine Network (1999), 10(3),

365 - 372

CODEN: ECYNEJ; ISSN: 1148-5493

PUBLISHER: John Libbey Eurotext

DOCUMENT TYPE: Journal LANGUAGE: English

Besides its role as a barrier against potential pathogens, intestinal flora is presumed to protect the host by priming the immunol. defense mechanisms. this respect, the influence of intestinal flora on macrophage precursors was examined, and its modulating effect was compared on LPS-induced cytokine production by macrophages derived from bone marrow and spleen precursors (BMDM and SDM resp.). The regulation of IL-1, IL-6, TNF- α and IL-12 production in macrophages from germfree and from three groups of flora-associated mice, conventional, conventionalized and E. coli-mono-associated mice, was investigated. The whole flora inhibited IL-1, TNF- α and IL-12 secretion by BMDM, whereas it had a stimulatory effect on IL-12 secretion by SDM. Implantation of $\mathbb E$. coli alone enhanced cytokine secretion by BMDM but had a more limited effect than whole flora on SDM, enhancing only TNF-lphaand IL-12 secretion. Study of expression of mRNA showed a correlation with protein secretion for IL-6 but not for TNF- α and IL-1. IL-12 enhancement in BMDM seemed to be dependent on regulation of p35 mRNA expression while it was correlated to increased p40 mRNA expression in SDM. The results demonstrated that intestinal flora modulated bone marrow and spleen macrophage cytokine production in a differential manner and suggested a role for bacteria other than %. coli among the whole flora. The contrasting effects exerted by the intestinal flora on bone marrow and spleen precursors are an interesting observation in view of the different functions of these organs in immunity. The finding that intestinal flora enhanced IL-12 production in spleen is also potentially important since this cytokine is implicated in the determination of the relative levels of Th1 and Th2 responses and plays a pivotal role in host defense against intracellular microorganisms. OS.CITING REF COUNT: 14THERE ARE 14 CAPLUS RECORDS THAT CITE THIS

RECORD (14 CITINGS)

REFERENCE COUNT: 50 THERE ARE 50 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 18 OF 21 MEDLINE on STN

ACCESSION NUMBER: 1998017378 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 9376506

TITLE: Construction of a Salmonalla typhimurium vaccine

strain expressing Vibrio cholera CT-B and

 ${\tt LPS-O}$ antigen.

AUTHOR: Chen D; Ma Q

CORPORATE SOURCE: Institute of Biotechnology, Academy of Military Medical

Sciences, Beijing, China.

SOURCE: Chinese journal of biotechnology, (1997) Vol. 13,

No. 1, pp. 43-50.

Journal code: 9100855. ISSN: 1042-749X.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199711

ENTRY DATE: Entered STN: 24 Dec 1997

Last Updated on STN: 24 Dec 1997

Entered Medline: 7 Nov 1997

The genes encoding V. cholera CT-B and LFS-O antigens were simultaneously inserted into the vector plasmid pYA248. The resulting recombinant plasmid pMG306 was transformed into the delta cya delta crp delta asd attenuated S. typhimurium vaccine strain x4072, and the live vaccine strain x4072 (pMG306) was constructed. This vaccine strain could secrete a specific CT-B antigen. Meanwhile, LPS-O antigens of both V. cholera and S. typhimurium were expressed on the cell surface. Mouse intraperitoneal immunization and subsequent challenge trial indicated that x4072 (pMG306) provided good protection against virulent V. cholera. This study has laid the foundation for the development of a new cholera-typhoid bivalent live oral vaccine.

L21 ANSWER 19 OF 21 MEDLINE on STN

ACCESSION NUMBER: 1997065056 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 8908606

TITLE: Lipopolysaccharides of Escherichia coli K12 strains that

express cloned genes for the Ogawa and Inaba

antigens of Vibrio cholerae O1: identification of $\ensuremath{\mathtt{O}}$

-antigenic factors.

AUTHOR: Hisatsune K; Kondo S; Iguchi T; Ito T; Hiramatsu K

CORPORATE SOURCE: Department of Microbiology, School of Pharmaceutical

Sciences, Josai University, Saitama, Japan.

SOURCE: Microbiology and immunology, (1996) Vol. 40, No.

9, pp. 621-6.

Journal code: 7703966. ISSN: 0385-5600.

PUB. COUNTRY: Japan

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199702

ENTRY DATE: Entered STN: 6 Mar 1997

Last Updated on STN: 6 Mar 1997 Entered Medline: 27 Feb 1997

AB Structural and serological studies were performed with the lipopolysaccharide (LPS) expressed by Escherichia coli K12 strains Number 30 and Number 64, into which cosmid clones derived from Vibrio cholerae O1 NIH 41 (Ogawa) and NIH 35A3 (Inaba) had been introduced, respectively. The two recombinant strains, Number 30 (Ogawa) and Number 64 (Inaba), produced LPS that included, in common, the O-polysaccharide chain composed of an alpha (1-->2)-linked N-(3-

deoxy-L-glycero-tetronyl)-D-perosamine (4-amino-4,6-dideoxy-D-manno-pyranose) homopolymer attached to the core oligosaccharide of the LFS of E. coli K12. Structural analysis revealed the presence of N-(3-deoxy-L-glycero-tetronyl)-2-O-methyl-D-perosamine at the non-reducing terminus of the O-polysaccharide chain of LPS from Number 30 (Ogawa) but not from Number 64 (Inaba). Serological analysis revealed that Number 30 (Ogawa) and Number 64 (Inaba) LPS were found to share the group antigen factor A of V. cholerae O1. They were distinguished by presence of the Ogawa antigen factor B [co-existing with relatively small amounts of the Inaba antigen factor (c)] in the former LPS and the Inaba antigen factor C in the latter LPS. It appears, therefore, that Number 30 (Ogawa) and Number 64 (Inaba) have Q- antigenic structures that are fully consistent with the AB(c) structure for the Ogawa and the AC structure for the Inaba O-forms of V. cholerae O1, respectively. Thus, the present study clearly confirmed our previous finding that the Ogawa antigenic factor B is substantially related to the 2-0-methyl group at the non-reducing terminus of the alpha (1-->2)-linked N-(3-deoxy-L-qlycero-tetronyl)-D-perosamine homopolymer that forms the O-polysaccharide chain of LPS of V. cholerae O1 (Ogawa).

L21 ANSWER 20 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1990:31565 HCAPLUS Full-text

DOCUMENT NUMBER: 112:31565

ORIGINAL REFERENCE NO.: 112:5365a,5368a

TITLE: Regulated expression of

proenkephalin A in normal lymphocytes

AUTHOR(S): Rosen, Haim; Behar, Oded; Abramsky, Oded; Ovadia, Haim

CORPORATE SOURCE: Fac. Med., Hebrew Univ., Jerusalem, Israel SOURCE: Journal of Immunology (1989), 143(11),

3703-7

CODEN: JOIMA3; ISSN: 0022-1767

DOCUMENT TYPE: Journal LANGUAGE: English

The expression of proenkephalin A (PEA), a neuropeptide-encoding gene, was examined in normal rat lymphocytes. With the use of Northern blot hybridization anal. of total RNA, PEA mRNA was found in normal cells derived from spleen, lymph nodes, and bone marrow. Cell sorting of the 2 main fractions of B and T cells derived from the spleen revealed that PEA is expressed in normal B cells (sIg+). The expression of PEA mRNA was markedly enhanced after a short incubation (3 h) of cells with LPS or Salmonella typhimurium. This was not the case when these cells were incubated with Con A during the same period of time; whereas, in thymocytes the presence of PEA mRNA was exclusively dependent upon mitogenic stimulus (Con A) and could be detected after 24 h of in vitro incubation. Exts. of cells were also found to contain immune reactive enkephalins, indicating that the PEA mRNA is translated. These results support the concept that neuropeptides, such as enkephalins, have a role in the modulation of the immune response and may participate in the bidirectional communication between the nervous and immune systems. OS.CITING REF COUNT: 29 THERE ARE 29 CAPLUS RECORDS THAT CITE THIS RECORD (29 CITINGS)

L21 ANSWER 21 OF 21 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1988:490879 HCAPLUS Full-text

DOCUMENT NUMBER: 109:90879

ORIGINAL REFERENCE NO.: 109:15151a,15154a

TITLE: Regulation of MHC expression in

vivo. Bacterial lipopolysaccharide induces class I

and II MHC products in mouse tissues by a T

cell-independent, cyclosporine-sensitive mechanism

AUTHOR(S): Jephthah-Ochola, Jessica; Urmson, Joan; Farkas, Susan;

Halloran, Philip F.

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792-800

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The effect of injections of bacterial LPS on the expression of class I and II products of the MHC in mouse tissues was investigated. In mice given two i.p. injections of LPS from Escherichia coli or Salmonalla minnesota, there were increases in class I and II MHC products in kidney, livex, heart, lung, and pancreas. Focusing on the changes in kidney, it was demonstrated that the increase in MHC expression occurred in tubules and, in the case of class I, in glomeruli. LPS treatment also increased the deposition of Ig in glomeruli. Time course studies indicated that increased class I empression could be induced by a single LPS injection, whereas class II induction required a second injection. The induction was influenced by the LPS sensitivity of the mouse strain, being much greater in LPS-sensitive C3H/HeSn mice than in LPS-resistant C3H/HeJ mice. T cells were not required. Nevertheless, the effect of LDS was inhibitable by cyclosporine and by a mAb against interferon- γ (IFN- γ) indicating that IFN- γ was required for the THERE ARE 22 CAPLUS RECORDS THAT CITE MHC induction. OS.CITING REF COUNT: 22 THIS

RECORD (22 CITINGS)

SEARCH HISTORY

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L2

L3

L4

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FILE 'HCAPLUS' ENTERED AT 17:22:48 ON 21 SEP 2009

E CURTISS III ROY/AU

E CURTISS ROY/AU

216 SEA ABB=ON ("CURTISS ROY"/AU OR "CURTISS ROY III"/AU) L1

9 SEA ABB=ON L1 AND ?LIVE?(W)?VACCINE?

2 SEA ABB=ON L2 AND ?CROSS?(W)?PROTECT?

ANALYZE L3 1-2 CT : 9 TERMS

182203 SEA ABB=ON ?SALMONELLA? OR E(W)?COLI? L5

L6 33841 SEA ABB=ON L5 AND (?LIVE? OR ?ATTENUAT? OR ?DERIV?)

L7 744 SEA ABB=ON L6 AND ?REGULAT?(5A)?EXPRES?

744 SEA ABB=ON L7 AND (?REGULAT? OR ?PROMOT? OR FUR OR ?INTEST? L8 OR ?EXPRES?)

L9 189 SEA ABB=ON L8 AND (?CARBOHYDRAT?(W)?ANTIGEN? OR IN(W)?VIVO? OR ?CROSS?(W)?PROTECT? OR ?IMMUN?)

FILE 'REGISTRY' ENTERED AT 17:30:20 ON 21 SEP 2009

E LPS O/CN

E LPS-O/CN

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FILE 'HCAPLUS' ENTERED AT 17:30:47 ON 21 SEP 2009
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42 SEA ABB=ON L8 AND LPS
L10
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L11 21 SEA ABB=ON L9 AND LPS

1 SEA ABB=ON L11 AND O(W)?ANTIGEN? L12

O SEA ABB=ON L11 AND PMI L13

1 SEA ABB=ON L9 AND PMI L14

L15

2 SEA ABB=ON L12 OR L14 22 SEA ABB=ON L11 OR L12 OR L14 L16

L17 18 SEA ABB=ON L16 AND (PRD<20070418 OR PD<20070418)

FILE 'MEDLINE, BIOSIS, EMBASE, DRUGU' ENTERED AT 17:34:12 ON 21 SEP 2009

114 SEA ABB=ON L17 L18

114 SEA ABB=ON L18 AND (LPS OR PMI) L19

3 SEA ABB=ON L19 AND O(W) ?ANTIGEN? L20

FILE 'HCAPLUS, MEDLINE' ENTERED AT 17:35:58 ON 21 SEP 2009 21 DUP REMOV L17 L20 (0 DUPLICATES REMOVED) L21

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